

We claim:

1. A method of determining enzyme activity comprising:

contacting a compound selected from the group consisting of enzymes,  
enzyme fragments and abzymes with a labeled substrate to form a  
differentially-charged product;

selectively coupling either the substrate or the differentially-charged product to  
an ion-exchange resin thereby substantially separating the substrate from the  
differentially-charged product; and

determining the amount of substrate remaining or differentially-charged  
product formed using a measuring means.

2. A method of determining enzyme activity comprising:

contacting a compound selected from the group consisting of enzymes,  
enzyme fragments and abzymes with a labeled substrate thereby effecting the  
conversion of the substrate to a differentially-charged product;

stopping the conversion before all of the substrate present has been converted  
to the differentially-charged product;

selectively coupling either the substrate or the differentially-charged product to  
an ion-exchange resin thereby substantially separating the substrate from the  
differentially-charged product in a single step; and

determining the amount of substrate remaining or differentially-charged  
product formed using a measuring means;

wherein the stopping step and coupling step are carried out concurrently or  
sequentially.

3. The method of claim 1 or 2 wherein the product is bound to the resin.

4. The method of claim 1 or 2 wherein the substrate is bound to the resin.

5. The method of claim 1 or 2 wherein the product or substrate measured is  
coupled to the resin

6. The method of claim 1 or 2 wherein the product or substrate measured is in solution
7. The method of claim 1 or 2 wherein said enzyme is a kinase.
8. The method of claim 1 or 2 wherein said method is conducted in a multiple-well format.
9. The method of claim 8 wherein the format comprises at least about 96 wells.
10. The method of claim 8 wherein said format is automated.
11. The method of claim 1 or 2 wherein said high-throughput format is conducted on a microchip.
12. The method of claim 1 or 2 wherein said enzyme is selected from the group consisting of GFAT, Nitric Oxide Synthase, Methionine Aminopeptidase, Asn Syn, PFK, p38, I-kappa kinase 1, I-kappa kinase 2, TBK1, MAPKAP 2, GTase, , OGTase, and Cyclooxygenase.
13. A method for identifying a molecule, compound, or composition that affects the activity of an enzyme, comprising:  
contacting the enzyme with a test sample comprising a molecule, compound, or composition;  
contacting the enzyme with a labeled substrate to form a differentially-charged product;

selectively coupling either the substrate or the differentially-charged product to an ion-exchange resin thereby substantially separating the substrate from the differentially-charged product;

5 determining the amount of substrate remaining or differentially-charged product formed using a measuring means; and

comparing the amount of substrate remaining or differentially-charged product formed with a control.

10      14.      The method of claim 13 wherein said enzyme is selected from the group consisting of GFAT, Nitric Oxide Synthase, Methionine Aminopeptidase, Asn Syn, PFK, p38, I-kappa kinase 1, I-kappa kinase 2, TBK1, MAPKAP 2, GTase, , OGTase, and Cyclooxygenase.

15     15.     The method of claim 13 wherein the control is an izozyme and the method is used to identifying a compound or composition that preferentially or specifically effects an enzyme over its isozyme.

16. A method of determining bi-functional enzyme activity comprising:

contacting an enzyme with a first labeled substrate to form a first differentially-charged product;

contacting the enzyme with a second labeled substrate to form a second differentially-charged product;

selectively coupling to an ion-exchange resin a member selected from the group consisting of the first substrate, the second substrate, the first product, and the second product, thereby substantially separating said member from the remaining members of the group; and

determining the amount of said member using a measuring means.

17. A method of determining bi-functional enzyme activity comprising:  
 35 contacting an enzyme with a first labeled substrate to form a first  
 differentially-charged product;  
 contacting the enzyme with a second labeled substrate to form a second  
 differentially-charged product;

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comparing the amount of substrate remaining or differentially-charged product formed from the plurality of enzymes.

21. A method of evaluating the selective coupling of an enzyme and a reactant comprising  
contacting a compound with an enzyme with a plurality of labeled substrates to  
form differentially-charged products;  
5 selectively coupling either the substrates or the differentially-charged products to an ion-exchange resin thereby substantially separating the substrates from the differentially-charged products;  
10 determining the amount of substrate remaining or differentially-charged products formed using a measuring means; and  
comparing the amount of substrate remaining or differentially-charged  
products formed from the plurality of substrates.  
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22. A kit for determining enzyme activity wherein said kit comprises at least three members of the group consisting of: An enzyme, a labeled ligand, a buffer solution, an ion-exchange resin, and a stop-buffer solution
- 20 23. A kit of claim 22 for determining enzyme activity wherein said kit comprises at least three members of the group consisting of: An enzyme, a labeled ligand, a buffer solution, an ion-exchange resin, and a stop-buffer solution
24. A kit of claim 23 for determining enzyme activity wherein said kit comprises  
25 at least three members of the group consisting of: An enzyme, a labeled ligand, a buffer solution, an ion-exchange resin, and a stop-buffer solution
25. A kit for determining enzyme activity comprising an enzyme, a labeled ligand, a buffer solution, an ion-exchange resin, and a stop-buffer solution  
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26. A compound discovered using the method of claims 1 or 2.

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